

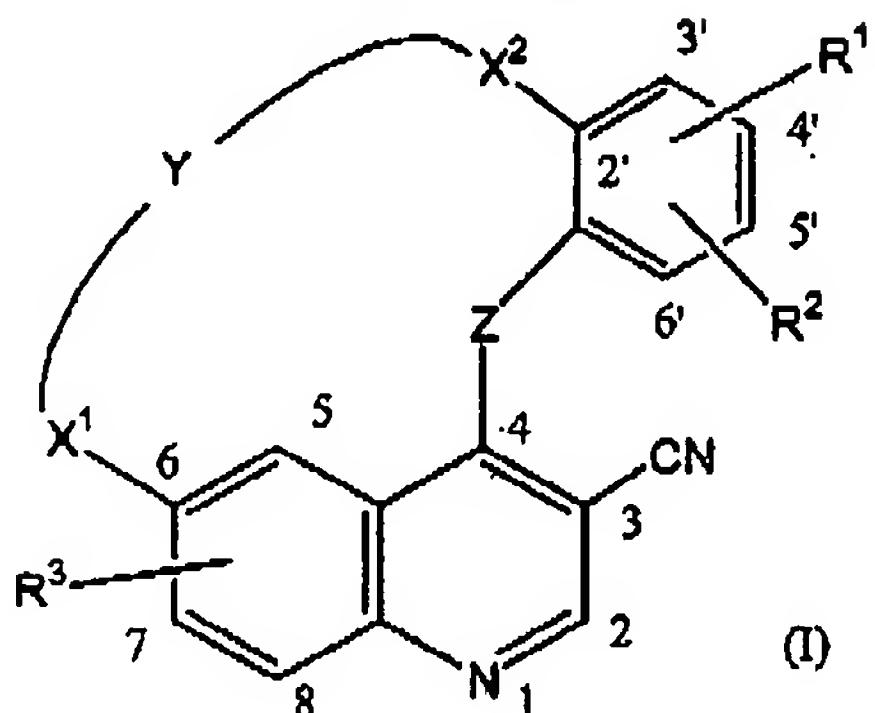
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PRD2168USPCT

Listing of Claims:

This listing of claims replaces all prior versions, and listings, of claims in the captioned application.

1. **(currently amended)** A compound having the formula



the N-oxide forms, the pharmaceutically acceptable addition salts and the stereochemically isomeric forms thereof, wherein

Z represents NH;

Y represents -C₁₋₅alkylene-NR¹²-C₁₋₅alkylene;

X¹ represents O;

X² represents a direct bond;

R¹ represents hydrogen, cyano, halo, hydroxy, formyl, C₁₋₆alkoxy-, C₁₋₆alkyl-,

C₁₋₆alkoxy- substituted with halo,

C₁₋₄alkyl substituted with one or where possible two or more substituents selected from hydroxy or halo;

R² represents hydrogen, cyano, halo, hydroxy, hydroxycarbonyl-, Het¹⁶-carbonyl-,

C₁₋₄alkyloxycarbonyl-, C₁₋₄alkylcarbonyl-, aminocarbonyl-,

mono- or di(C₁₋₄alkyl)aminocarbonyl-, Het¹, formyl, C₁₋₄alkyl-, C₂₋₆alkynyl-,

C₃₋₆cycloalkyl-, C₃₋₆cycloalkyloxy-, C₁₋₆alkoxy-, Ar⁵, Ar¹-oxy-, dihydroxyborane ,

C₁₋₆alkoxy- substituted with halo,

C₁₋₄alkyl substituted with one or where possible two or more substituents selected from halo, hydroxy or NR⁴R⁵,

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$C_{1-4}alkylcarbonyl$ - wherein said $C_{1-4}alkyl$ is optionally substituted with one or where possible two or more substituents selected from hydroxy or

$C_{1-4}alkyl-oxy$ -;

R^3 represents hydrogen, hydroxy, Ar^3 -oxy, $Ar^4-C_{1-4}alkyloxy$ -, $C_{1-4}alkyloxy$ -,

$C_{2-4}alkenyloxy$ - optionally substituted with Het^{12} or R^3 represents $C_{1-4}alkyloxy$ substituted with one or where possible two or more substituents selected from

$C_{1-4}alkyloxy$ -, hydroxy, halo, Het^2 -, $-NR^6R^7$, -carbonyl-, NR^8R^9 or Het^3 -carbonyl-;

R^4 and R^5 are each independently selected from hydrogen or $C_{1-4}alkyl$;

R^6 and R^7 are each independently selected from hydrogen, $C_{1-4}alkyl$, Het^8 , aminosulfonyl-, mono- or di($C_{1-4}alkyl$)-aminosulfonyl, hydroxy- $C_{1-4}alkyl$ -,

$C_{1-4}alkyl-oxy-C_{1-4}alkyl$ -, hydroxycarbonyl- $C_{1-4}alkyl$ -, $C_{3-6}cycloalkyl$, Het^9 -carbonyl- $C_{1-4}alkyl$ -, Het^{10} -carbonyl-, polyhydroxy- $C_{1-4}alkyl$ -, Het^{11} - $C_{1-4}alkyl$ - or $Ar^2-C_{1-4}alkyl$ -;

R^8 and R^9 are each independently selected from hydrogen, $C_{1-4}alkyl$, $C_{3-6}cycloalkyl$, Het^4 , hydroxy- $C_{1-4}alkyl$ -, $C_{1-4}alkyloxyC_{1-4}alkyl$ - or polyhydroxy- $C_{1-4}alkyl$ -;

R^{12} represents hydrogen, $C_{1-4}alkyl$, Het^{13} , $Het^{14}-C_{1-4}alkyl$ - or phenyl optionally substituted with one or where possible two or more substituents selected from hydrogen, hydroxy, amino or $C_{1-4}alkyloxy$ -;

Het^1 represents a heterocycle selected from piperidinyl, morpholinyl, piperazinyl, furanyl, pyrazolyl, dioxolanyl, thiazolyl, oxazolyl, imidazolyl, isoxazolyl, oxadiazolyl, pyridinyl or pyrrolidinyl wherein said Het^1 is optionally substituted amino, $C_{1-4}alkyl$, hydroxy- $C_{1-4}alkyl$ -, phenyl, phenyl- $C_{1-4}alkyl$ -,

$C_{1-4}alkyl-oxy-C_{1-4}alkyl$ - mono- or di($C_{1-4}alkyl$)amino- or amino-carbonyl-;

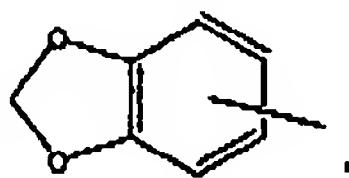
Het^2 represents a heterocycle selected from morpholinyl, piperazinyl, piperidinyl, pyrrolidinyl, thiomorpholinyl or dithianyl wherein said Het^2 is optionally substituted with one or where possible two or more substituents selected from hydroxy, halo, amino, $C_{1-4}alkyl$ -, hydroxy- $C_{1-4}alkyl$ -, $C_{1-4}alkyl-oxy-C_{1-4}alkyl$ -, hydroxy- $C_{1-4}alkyl-oxy-C_{1-4}alkyl$ -, mono- or di($C_{1-4}alkyl$)amino-, mono- or di($C_{1-4}alkyl$)amino-C₁₋₄alkyl-, amino- $C_{1-4}alkyl$ -, mono- or di($C_{1-4}alkyl$)amino-sulfonyl-, aminosulfonyl-;

Het^3 , Het^4 and Het^8 each independently represent a heterocycle selected from morpholinyl, piperazinyl, piperidinyl, furanyl, pyrazolyl, dioxolanyl, thiazolyl, oxazolyl, imidazolyl, isoxazolyl, oxadiazolyl, pyridinyl or pyrrolidinyl wherein said Het^3 , Het^4 or Het^8 is optionally substituted with one or where possible two or more substituents selected from hydroxy-, amino-, $C_{1-4}alkyl$ -, $C_{3-6}cycloalkyl-C_{1-4}alkyl$ -, aminosulfonyl-, mono- or di($C_{1-4}alkyl$)aminosulfonyl or amino- $C_{1-4}alkyl$ -;

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Het⁹ and Het¹⁰ each independently represent a heterocycle selected from furanyl, piperidinyl, morpholinyl, piperazinyl, pyrazolyl, dioxolanyl, thiazolyl, oxazolyl, imidazolyl, isoxazolyl, oxadiazolyl, pyridinyl or pyrrolidinyl wherein said Het⁹ or Het¹⁰ is optionally substituted C₁₋₄alkyl, C₃₋₆cycloalkyl-C₁₋₄alkyl- or amino-C₁₋₄alkyl-;



Het¹¹ represents a heterocycle selected from indolyl or

Het¹² represents a heterocycle selected from morpholinyl, piperazinyl, piperidinyl, pyrrolidinyl, thiomorpholinyl or dithianyl wherein said Het¹² is optionally substituted with one or where possible two or more substituents selected from hydroxy, halo, amino, C₁₋₄alkyl-, hydroxy-C₁₋₄alkyl-, C₁₋₄alkyl-oxy-C₁₋₄alkyl-, hydroxy-C₁₋₄alkyl-oxy-C₁₋₄alkyl-, mono- or di(C₁₋₄alkyl)amino- or mono- or di(C₁₋₄alkyl)amino-C₁₋₄alkyl-;

Het¹³ represent a heterocycle selected from pyrrolidinyl or piperidinyl wherein said Het¹³ is optionally substituted with one or where possible two or more substituents selected from C₁₋₄alkyl, C₃₋₆cycloalkyl, hydroxy-C₁₋₄alkyl-, C₁₋₄alkyloxyC₁₋₄alkyl or polyhydroxy-C₁₋₄alkyl-;

Het¹⁴ represent a heterocycle selected from morpholinyl, pyrrolidinyl, piperazinyl or piperidinyl wherein said Het¹⁴ is optionally substituted with one or where possible two or more substituents selected from C₁₋₄alkyl, C₃₋₆cycloalkyl, hydroxy-C₁₋₄alkyl-, C₁₋₄alkyloxyC₁₋₄alkyl or polyhydroxy-C₁₋₄alkyl-;

Het¹⁶ represent a heterocycle selected from morpholinyl, pyrrolidinyl, piperazinyl, 1,3,2-dioxaborolane or piperidinyl wherein said heterocycle is optionally substituted with one or more substituents selected from C₁₋₄alkyl; and

Ar¹, Ar², Ar³, Ar⁴ and Ar⁵ each independently represent phenyl optionally substituted with cyano, C₁₋₄alkylsulfonyl-, C₁₋₄alkylsulfonylamino-, aminosulfonylamino-, hydroxy-C₁₋₄alkyl, aminosulfonyl-, hydroxy-, C₁₋₄alkyloxy- or C₁₋₄alkyl.

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2. (currently amended) A compound according to claim 1 wherein;

R¹ represents halo;

R² represents hydrogen, cyano, halo, hydroxy, hydroxycarbonyl-, C₁₋₄alkyloxycarbonyl-, Het¹⁶-carbonyl-, C₂₋₆alkynyl-, Ar⁵ or Het¹;

~~In a further embodiment R³ represents hydrogen, cyano, halo, hydroxy, C₂₋₆alkynyl or Het¹;~~

R³ represents hydrogen, hydroxy, C₁₋₄alkyloxy-, Ar⁴-C₁₋₄alkyloxy or R³ represents

C₁₋₄alkyloxy substituted with one or where possible two or more substituents selected from C₁₋₄alkyloxy- or Het²-;

R¹² represents Het¹⁴-C₁₋₄alkyl, in particular morpholinyl-C₁₋₄alkyl;

Het¹ represents thiazolyl optionally substituted amino, C₁₋₄alkyl, hydroxy-C₁₋₄alkyl-, phenyl, phenyl-C₁₋₄alkyl-, C₁₋₄alkyl-oxy-C₁₋₄alkyl- mono- or di(C₁₋₄alkyl)amino- or amino-carbonyl;

Het² represents a heterocycle selected from morpholinyl, piperazinyl, piperidinyl or pyrrolidinyl wherein said Het² is optionally substituted with one or where possible two or more substituents selected from hydroxy, amino or C₁₋₄alkyl-;

Het¹⁴ represents a heterocycle selected from morpholinyl, piperazinyl, piperidinyl or pyrrolidinyl wherein said Het¹⁴ is optionally substituted with one or where possible two or more substituents selected from hydroxy, amino or C₁₋₄alkyl-;

Het¹⁶ represents a heterocycle selected from piperidinyl, morpholinyl or pyrrolidinyl;

Ar⁴ represents phenyl optionally substituted with cyano, hydroxy-, C₁₋₄alkyloxy or C₁₋₄alkyl; and

Ar⁵ represents phenyl optionally substituted with cyano, hydroxy, C₁₋₄alkyloxy or C₁₋₄alkyl.

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3. (previously presented) A compound according to claim 1 wherein:
R¹ represents hydrogen or halo;
R² represents hydrogen, cyano, halo, hydroxycarbonyl-, C₁₋₄alkyloxycarbonyl-, Het¹⁶-carbonyl- or Ar⁵;
R³ represents hydrogen, hydroxy, C₁₋₄alkyloxy-, Ar⁴-C₁₋₄alkyloxy or R³ represents C₁₋₄alkyloxy substituted with one or where possible two or more substituents selected from C₁₋₄alkyloxy- or Het²;
Het² represents a heterocycle selected from morpholinyl, piperazinyl, piperidinyl or pyrrolidinyl wherein said Het² is optionally substituted with one or where possible two or more substituents selected from hydroxy, amino or C₁₋₄alkyl-;
Het¹⁴ represents morpholinyl;
Het¹⁶ represents a heterocycle selected from morpholinyl or pyrrolidinyl;
Ar⁴ represents phenyl; and
Ar⁵ represents phenyl optionally substituted with cyano.

4. (currently amended) A compound according to claim 1, wherein the R¹ substituent is at position 4', the R² substituent is at position 5' and the R³ substituent is at position 7 of the structure of formula (I).

5.-7. (Cancelled)

8. (previously presented) A pharmaceutical composition comprising a compound as described in Claim 1 and a pharmaceutically acceptable carrier.

9.-12 (cancelled)